Docket No. ARS-113 Patent Application 10/540845 JC17 Rec'd PCT/PTO 27 JUN 2005

In the Claims

1-41 (Canceled).

42 (New). A composition of matter comprising:

- a) an isolated polypeptide having Preadipocyte-Factor-1-like activity selected from the group consisting of: 1) the amino acid sequence recited in SEQ ID NO: 2; 2) the mature form of the polypeptide whose sequence is recited in SEQ ID NO: 2 (SEQ ID NO:4); 3) active variants of the amino acid sequence of SEQ ID NO: 2, wherein any amino acid specified in the chosen sequence is non-conservatively substituted, provided that no more than 15% of the amino acid residues in the sequence are so changed; and 4) the active fragment, precursor, salt, or derivative of the amino acid sequences given in 1), 2), or 3);
- b) an isolated polypeptide that is a naturally occurring allelic variant of the sequence given by SEQ ID NO: 2 or SEQ ID NO:4;
- c) an isolated polypeptide that is a naturally occurring allelic variant of the sequence given by SEQ ID NO: 2 or SEQ ID NO:4, wherein the variant is the translation of a single nucleotide polymorphism;
- d) a polypeptide as set forth in a) or b) or c), wherein the polypeptide binds specifically an antibody or a binding protein generated against SEQ ID NO: 2 or SEQ ID NO:4 or a fragment thereof;
 - e) a fusion protein comprising a polypeptide as set forth in a) or b) or c) or d);
- f) an antagonist of a polypeptide as set forth in a) or b) or c) or d), wherein said antagonist comprises an amino acid sequence resulting from the non-conservative substitution, the deletion or both the non-conservative substitution and deletion of one or more residues into the corresponding polypeptide;
 - g) a ligand which binds specifically to a polypeptide a) or b) or c) or d);
- h) a polypeptide a) or b) or c) or d) or e), wherein said polypeptides are in the form of active conjugates or complexes with a molecule chosen from radioactive labels, fluorescent labels, biotin, or cytotoxic agents;

- i) a peptide mimetic designed on the sequence or the structure or the sequence and structure of a polypeptide as set forth in a) or b) or c) or d);
- j) an isolated nucleic acid encoding for an isolated polypeptide selected from the group consisting of:
 - 1) polypeptides as set forth in a) or b) or c) or d);
 - 2) a fusion protein comprising a polypeptide as set forth in a) or b) or c) or d); or
 - an antagonist of a polypeptide as set forth in a) or b) or c) or d), wherein said antagonist comprises an amino acid sequence resulting from the non-conservative substitution, the deletion or both the non-conservative substitution and deletion of one or more residues into the corresponding polypeptide;
- k) an isolated nucleic acid sequence comprising SEQ ID NO: 1; nucleotides 122-1180 of SEQ ID NO: 1; or a complement of either sequence;
 - 1) a purified nucleic acid which:
 - 1) hybridizes under high stringency conditions; or
 - exhibits at least about 85% identity over a stretch of at least about 30 nucleotides with a nucleic acid selected from the group consisting of SEQ ID NO: 1; nucleotides 122-1180 of SEQ ID NO: 1; or a complement of either sequence;
 - m) a vector comprising a nucleic acid as set forth in j) or k) or l);
 - n) a polypeptide encoded by the nucleic acid of j) or k) or l);
 - o) a host cell comprising a vector or nucleic acid as set forth in j) or k) or l) or m);
- p) a transgenic animal cell comprising a vector or nucleic acid as set forth in j) or k) or l) or m) and having enhanced or reduced expression levels of a polypeptide as set forth in a) or b) or c) or d);
- q) a transgenic non-human animal that has been transformed to have enhanced or reduced expression levels of a polypeptide as set forth in a) or b) or c) or d);
- r) a compound that enhances the expression level of a polypeptide as set forth in a) or b) or c) or d) in a cell or animal; or

- s) a compound that reduces the expression level of a polypeptide as set forth in a) or b) or c) or d) in a cell or animal.
- 43. (New) The composition of matter according to claim 42, wherein said composition of matter comprises a polypeptide, peptide mimetic, nucleic acid, cell, or compound that enhances or reduces the expression of a polypeptide and a pharmaceutically acceptable carrier.
- 44. (New) The composition of matter according to claim 42, wherein the fusion protein further comprises one or more amino acid sequence selected from the protein sequences: membrane-bound protein, immunoglobulin constant region, multimerization domains, extracellular proteins, signal peptide-containing proteins, or export signal-containing proteins.
- 45. (New) The composition of matter according to claim 42, wherein the ligand antagonizes or inhibits the Preadipocyte-Factor-1-like activity of a polypeptide.
- 46. (New) The composition of matter according to claim 45, wherein the ligand is a monoclonal antibody, a polyclonal antibody, a humanized antibody, an antigen binding fragment, or the extracellular domain of a membrane-bound protein.
- 47. (New) The composition of matter according to claim 42, wherein said vector comprises a nucleic acid molecule that is operatively linked to expression control sequences allowing expression in prokaryotic or eukaryotic host cells of the encoded polypeptide.
- 48. (New) A method for determining the activity and/or the presence of a Preadipocyte-Factor-1-like polypeptide in a sample comprising:
 - a) providing a protein-containing sample;
- b) contacting said sample with a ligand that specifically binds to a polypeptide comprising:

- an isolated polypeptide having Preadipocyte-Factor-1-like activity selected from the group consisting of: i) the amino acid sequence recited in SEQ ID NO: 2; ii) the mature form of the polypeptide whose sequence is recited in SEQ ID NO: 2 (SEQ ID NO:4); iii) active variants of the amino acid sequence of SEQ ID NO: 2, wherein any amino acid specified in the chosen sequence is non-conservatively substituted, provided that no more than 15% of the amino acid residues in the sequence are so changed; and iv) the active fragment, precursor, salt, or derivative of the amino acid sequences given in i), ii), or iii);
- 2) an isolated polypeptide that is a naturally occurring allelic variant of the sequence given by SEQ ID NO: 2 or SEQ ID NO:4;
- an isolated polypeptide that is a naturally occurring allelic variant of the sequence given by SEQ ID NO: 2 or SEQ ID NO:4, wherein the variant is the translation of a single nucleotide polymorphism; or
- a polypeptide as set forth in a) or b) or c), wherein the polypeptide binds specifically an antibody or a binding protein generated against SEQ ID NO: 2 or SEQ ID NO:4 or a fragment thereof; and
- c) determining the presence or said ligand bound to said polypeptide.
- 49. (New) A method of using the composition of claim 42 for a) producing cells capable of expressing a polypeptide; b) making a polypeptide; c) the treatment of a disease; d) the preparation of pharmaceutical compositions; e) the screening candidate compounds; f) identifying a candidate compound; or g) determining the presence or the amount of a transcript or of a nucleic acid.
- 50. (New) The method according to claim 49, wherein said method comprises genetically engineering cells with a vector or a nucleic acid comprising:
- a) an isolated nucleic acid encoding for an isolated polypeptide selected from the group consisting of:
 - 1) an isolated polypeptide having Preadipocyte-Factor-1-like activity selected

from the group consisting of: i) the amino acid sequence recited in SEQ ID NO: 2; ii) the mature form of the polypeptide whose sequence is recited in SEQ ID NO: 2 (SEQ ID NO:4); iii) active variants of the amino acid sequence of SEQ ID NO: 2, wherein any amino acid specified in the chosen sequence is non-conservatively substituted, provided that no more than 15% of the amino acid residues in the sequence are so changed; and iv) the active fragment, precursor, salt, or derivative of the amino acid sequences given in i), ii), or iii);

- 2) an isolated polypeptide that is a naturally occurring allelic variant of the sequence given by SEQ ID NO: 2 or SEQ ID NO:4;
- an isolated polypeptide that is a naturally occurring allelic variant of the sequence given by SEQ ID NO: 2 or SEQ ID NO:4, wherein the variant is the translation of a single nucleotide polymorphism;
- 4) a polypeptide as set forth in 1) or 2) or 3), wherein the polypeptide binds specifically an antibody or a binding protein generated against SEQ ID NO: 2 or SEQ ID NO:4 or a fragment thereof; or
- 5) a fusion protein comprising a polypeptide as set forth in 1) or 2) or 3) or 4);
- b) an isolated nucleic acid sequence comprising SEQ ID NO: 1; nucleotides 122-1180 of SEQ ID NO: 1; or a complement of either sequence;
 - c) a purified nucleic acid which:
 - 1) hybridizes under high stringency conditions; or
 - exhibits at least about 85% identity over a stretch of at least about 30 nucleotides with a nucleic acid selected from the group consisting of SEQ ID NO: 1; nucleotides 122-1180 of SEQ ID NO: 1; or a complement of either sequence; or
 - d) a vector comprising a nucleic acid as set forth in a) or b) or c).
- 51. (New) The method according to claim 49, wherein said method comprises a method for making a polypeptide comprising culturing a transformed host cell under conditions in which the

nucleic acid or vector is expressed, and recovering the polypeptide encoded by said nucleic acid or vector from the culture.

- 52. (New) The method according to claim 49, wherein said method comprises the treatment of a disease needing an increase in the Preadipocyte-Factor-1-like activity that comprises the administration of a therapeutically effective amount of a polypeptide, a peptide mimetic, a nucleic acid, a cell, or a compound as set forth in claim 42.
- 53. (New) The method according to claim 49, wherein said method of treatment comprises the administration of a therapeutically effective amount of an antagonist, a ligand, or of a compound as set forth in claim 42.
- 54. (New) The method according to claim 49, wherein said method for screening candidate compounds effective to treat a disease related to the Preadipocyte-Factor-1-like polypeptides comprises:
- a) contacting a cell, a transgenic animal cell, or a transgenic non-human animal having enhanced or reduced expression levels of the Preadipocyte-Factor-1-like polypeptide, with a candidate compound; and
 - b) determining the effect of the compound on the animal or on the cell.
- 55. (New) The method according to claim 49, wherein said method for identifying a candidate compound as an antagonist/inhibitor or agonist/activator of a Preadipocyte-Factor-1-like polypeptide comprises:
- a) contacting said polypeptide, said compound, and a mammalian cell or a mammalian cell membrane capable of binding the polypeptide; and
- b) measuring whether the molecule blocks or enhances the interaction of the polypeptide, or the response that results from such interaction, with the mammalian cell or the mammalian cell membrane.

- 56. (New). The method according to claim 49, wherein said method for determining the presence or the amount of a transcript or of a nucleic acid encoding a Preadipocyte-Factor-1-like polypeptide comprises:
 - a) providing a nucleic acids-containing sample;
 - b) contacting said sample with a nucleic acid as set forth in claim 42; and
 - c) determining the hybridization of said nucleic acid with a nucleic acid into the sample.